✓ATENT COOPERATION TREATY

	From the INTERNATIONAL BUREAU
PCT	То:
NOTIFICATION OF ELECTION	United States Patent and Trademark Office
(PCT Rule 61.2)	(Box PCT) Crystal Plaza 2 Washington, DC 20231 ÉTATS-UNIS D'AMÉRIQUE
Date of mailing (day/month/year) 09 July 1999 (09.07.99)	in its capacity as elected Office
International application No.	Applicant's or agent's file reference

International application No.
PCT/US98/25791

International filing date (day/month/year)
O4 December 1998 (04.12.98)

Applicant's or agent's file reference
LJIEM1100WO

Priority date (day/month/year)
O5 December 1997 (05.12.97)

Applicant
BOURDON, Mario, A. et al

1.	The designated Office is hereby notified of its election made:
	X in the demand filed with the International Preliminary Examining Authority on:
	04 June 1999 (04.06.99)
	in a notice effecting later election filed with the International Bureau on:
2.	The election X was
	was not
	made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland **Authorized officer**

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D.E.

PATENT COOPERATION TREATY

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INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

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LJIEM1100WO	FOR FURTHER ACTION	See Notifica	tion of Transmittal of International		
International application No.			xamination Report (Form PCT/IPEA/416)		
1	International filing date (day/m	, , , , , , , , , , , , , , , , , , , ,			
PCT/US98/25791	04 DECEMBER 1998		05 DECEMBER 1997		
International Patent Classification (IPC) of Please See Supplemental Sheet.	or national classification and IPC	!			
Applicant LA JOLLA INSTITUTE FOR EXPERII	MENTAL MEDICINE				
2. This REPORT consists of a to the This report is also accomplished and are the (see Rule 70.16 and Section 1).	total of sheets. panied by ANNEXES, i.e., sheete basis for this report and/or she ion 607 of the Administrative I	ccording to Ar ts of the descrip	tion, claims and/or drawings which have		
These annexes consist of a to	tal of sheets.				
3. This report contains indication	s relating to the following ite	ems:			
I Desire (A)	-				
	ı				
II Priority					
III Non-establishmen	t of report with regard to nov	elty, inventive	step or industrial applicability		
IV Lack of unity of invention					
V X Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement					
VI Certain documents of					
VII Certain defects in the international application					
VIII Certain observations on the international application					
Date of submission of the demand		£1-4: £	4		
or seemand	Date	f completion of	this report		
04 JUNE 1999	27	FEBRUARY 20	000		
Name and mailing address of the IPEA/U	S Aat	ized office-	10005 2010 201		
Commissioner of Patents and Tradema		ized officer	JOYCE BRIDGERS PARALEGAL SPECIALIST		
Box PCT Washington, D.C. 20231	su	SAN UNGAR	CHEMICAL MATRIX		
Facsimile No. (703) 305-3230	Teleph	one No. <i>(</i> 703)			

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application	No.
PCT/HS98/25791	

L Ba	asis of	the report		
1. This i	report ha	s been drawn on the	basis of Substitute sheets we this report as "originally file	hich have been furnished to the receiving Office in response to an invitation ed" and are not annexed to the report since they do not contain amendments):
	x		al application as origin	• •
	X		, pages <u>1-18</u>	
		-		, filed with the demand.
				_ , filed with the letter of
				, filed with the letter of
	X	the claims,	Nos. <u>1-29</u>	, as originally filed.
			Nos. NONE	, as amended under Article 19.
			Nos. NONE	, filed with the demand.
			Nos. NONE	, filed with the letter of
				, filed with the letter of
	x	the drawings,	sheets/ fig 1-3	, as originally filed.
	_		sheets /fig NONE	, filed with the demand.
			sheets/ fig NONE	, filed with the letter of
			sheets /fig	, filed with the letter of
	x	the claims,	Nos. none sheets/fig none	
3.	This to go	report has been eso beyond the disclo	stablished as if (some of) stablished as filed, as indicated	the amendments had not been made, since they have been considered in the Supplemental Box Additional observations below (Rule 70.2(c)).
4. Add		l observations, if	necessary:	
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INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

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V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. STATEMENT			
Novelty (N)	Claims	13-20, 23-25, 27-29	YES
	Claims	1-12, 21-22, 26	NO
Inventive Step (IS)	Claims	13-20, 23-25, 27-29	YES
	Claims	1-12, 21-22, 26	NO NO
Industrial Applicability (IA)	Claims	1-29	YES
	Claims	NONE	NO

2. CITATIONS AND EXPLANATIONS

Claims 1-2 lack novelty under PCT Article 33(2) as being anticipated by Borgstrom (1993). Borgstrom ('93) discloses a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of an agent that inhibits angiogenesis (Linomide) to a cellular system is interpreted to meet the claim limitation "inhibiting a host cell angiogenic effect."

Claims 1-5 and 26 lack novelty under PCT Article 33(2) as being anticipated by either of Ferrara or Borgstrom. Both references disclose a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of an agent that inhibits angiogenesis (anti-VEGF) is interpreted to meet the claim limitation "inhibiting a host cell angiogenic effect."

Claims 1-4 lack novelty under PCT Article 33(2) as being anticipated by either of DiPetro or Hu. Both references disclose a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of an agent that inhibits angiogenesis (TSP1, as in DiPetro, or IL-1 receptor antagonist, antiserum to IL-8, anti-TNF alpha, or monoclonal bFGF, as in Hu)is interpreted to meet the claim limitation "inhibiting a host cell angiogenic effect."

Claims 1-12, 21-22 and 26 lack novelty under PCT Article 33(2) as being anticipated by Lokeshwar. Lokeshwar discloses a method that is inherently the same as that claimed, a method for the inhibition of angiogenesis comprising inhibiting a host cell angiogenic effect that is the same as that claimed. The administration of the 5A1 monoclonal antibody to M-CSF-1, which inherently "inhibits a host cell angiogenic effect."

Claims 1-29 have industrial applicability as defined by PCT Article 33(4).

Claims 13-20, 23-25 and 27-29 meet the criteria set out in PC1 (Continued on Supplemental Sheet.)

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TATEMATIONAL PREDIVINARY EXAMINATION REPORT	PCT/US98/25791
Supplemental Box (To be used when the space in any of the preceding boxes is not sufficient))
Continuation of: Boxes I - VIII	Sheet 10
CLASSIFICATION: The International Patent Classification (IPC) and/or the National classific IPC(7): A61K 39/395, 48/00; C07K 16/18, 16/22, 16/24, 16/26, 16/28 and US C 388.85	ation are as listed below: Cl.: 424/130.1; 514/44; 530/387.1, 388.23,
V. 2. REASONED STATEMENTS - CITATIONS AND EXPLANATIONS (Confarticle 33(2)-(3), because the prior art does not teach or fairly suggest the claimed	inued): method of inhibiting angiogenesis.
LOKESHWAR, B.L. et al. Development and Characterization of Monoclonal And Stimulating Factor. The Journal of Immunology. 15 July 1988, Vol. 141, No. 2, p	ibodies to Murine Macrophage Colony- ages 483-488, see entire document.

INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/25791

IPC(6) US CL	SSIFICATION OF SUBJECT MATTER :A61K 39/395, 48/00; C07K 16/18, 16/22, 16/24, 16. :424/130.1; 514/44; 530/387.1, 388.23, 388.85 to International Patent Classification (IPC) or to both		d IPC		
	DS SEARCHED				
Minimum d	ocumentation searched (classification system followe	d by classification symbo	ls)		
	424/130.1; 514/44; 530/387.1, 388.23, 388.85				
Documenta	tion searched other than minimum documentation to the	extent that such documen	ats are included	in the fields searched	
Electronic d	lata base consulted during the international search (na	me of data base and, who	ere practicable,	search terms used)	
	N, REGISTRY, CAPLUS, MEDLINE, EMBASE, LIF of angiogenesis, macrophages, M-CSF, M-CSF-1	ESCI, BIOTECHDS, WIF	PIDS	·	
C. DOC	UMENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where ap	propriate, of the relevant	passages	Relevant to claim No.	
X	Database PREV, 1993: 44131748, BORGSTROM, P. et al. Inhibition of Angiogensis by the Quinoline-2-carboxamide Linomide. FASEB Journal. 28 March-April 1993, Vol. 7, No. 3-4, page A879, see entire document.			1-3, 13-14, 26	
X	FERRARA, Napoleone. The Role of Vascular Endothelial Growth Factor in Pathological Angiogenesis. Breast Cancer Research and Treatment. 1995, Vol. 36, pages 127-137, see entire document.			1-5, 13, 26	
Х	DiPIETRO, L. et al. Angiogenic Macrophages Produce the Angiogenic Inhibitor Thrombospondin 1. American Journal of Pathology. September 1993, Vol. 143, No. 3, pages 678-684, see entire document.				
X Furth	ner documents are listed in the continuation of Box C	. See patent fa	mily annex.		
'A' do					
	be of particular relevance rlier document published on or after the international filing date	"X" document of partic	cular relevance; the	claimed invention cannot be	
cit	*L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other				
special reason (as specified) "O" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art					
	·				
	Date of the actual completion of the international search 23 MARCH 1999 Date of mailing of the international search report 15 APR 1999				
Commission Box PCT	Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Authorized officer				
Washingto Facsimile N	n, D.C. 20231 No. (703) 305-3230	Telephone No. (703)	SON) 308-0196	for	
	•	(100)	,		

INTERNATIONAL SEARCH REPORT

International application No. PCT/US98/25791

		Relevant to claim No	
Category*	tegory* Citation of document, with indication, where appropriate, of the relevant passages		
x	BORGSTROM, P. et al. Complete Inhibition of Angiogenesis and Growth of Microtumors by Anti-Vascular Endothelial Growth Factor Neutralizing Antibody: Novel Concepts of Angiostatic Therapy from Intravital Videomicroscopy. Cancer Research. 01 September 1996, Vol. 56, pages 4032-4039, see entire document.	1-5, 26	
K	HU, D.E., et al. Inhibition of Angiogenesis in Rats by IL-1 Receptor Antagonist and Selected Cytokine Antibodies. Inflammation. 1994, Vol. 18, No.1, pages 45-58, see entire doiciumnet.	1-4	
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